# Serial#: 10/566,214 INVENTOR SEARCH

#### => FILE REG

FILE 'REGISTRY' ENTERED AT 18:31:38 ON 20 NOV 2009
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2009 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 19 NOV 2009 HIGHEST RN 1192927-49-3 DICTIONARY FILE UPDATES: 19 NOV 2009 HIGHEST RN 1192927-49-3

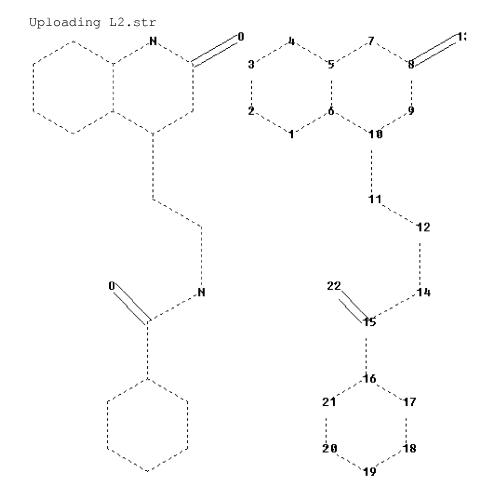
New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH June 26, 2009.

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

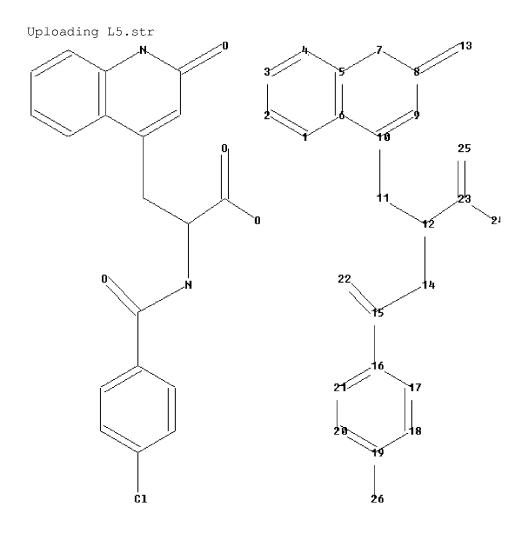
http://www.cas.org/support/stngen/stndoc/properties.html



chain nodes :
11 12 13 14 15 22
ring nodes :
1 2 3 4 5 6 7 8 9 10 16 17 18 19 20 21
chain bonds :
8-13 10-11 11-12 12-14 14-15 15-16 15-22
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-10 7-8 8-9 9-10 16-17 16-21 17-18 18-19
19-20 20-21
exact/norm bonds :
1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-10 7-8 8-9 8-13 9-10 10-11 11-12 12-14
14-15 15-16 15-22 16-17 16-21 17-18 18-19 19-20 20-21

## Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:CLASS

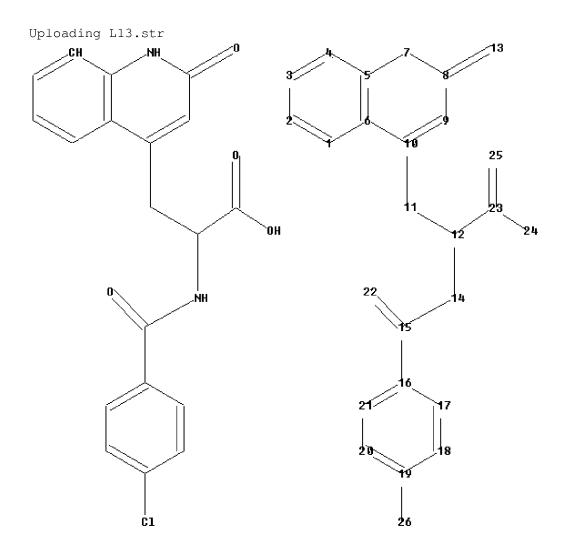


chain nodes :
11 12 13 14 15 22 23 24 25 26
ring nodes :

1 2 3 4 5 6 7 8 9 10 16 17 18 19 20 21 chain bonds:
8-13 10-11 11-12 12-14 12-23 14-15 15-16 15-22 19-26 23-24 23-25 ring bonds:
1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-10 7-8 8-9 9-10 16-17 16-21 17-18 18-19 19-20 20-21 exact/norm bonds:
5-7 6-10 7-8 8-9 8-13 9-10 12-14 14-15 15-22 23-24 23-25 exact bonds:
10-11 11-12 12-23 15-16 19-26 normalized bonds:
1-2 1-6 2-3 3-4 4-5 5-6 16-17 16-21 17-18 18-19 19-20 20-21

#### Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS



chain nodes : 11 12 13 14 15 22 23 24 25 26

ring nodes:
1 2 3 4 5 6 7 8 9 10 16 17 18 19 20 21
chain bonds:
8-13 10-11 11-12 12-14 12-23 14-15 15-16 15-22 19-26 23-24 23-25
ring bonds:
1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-10 7-8 8-9 9-10 16-17 16-21 17-18 18-19
19-20 20-21
exact/norm bonds:
5-7 6-10 7-8 8-9 8-13 9-10 12-14 14-15 15-22
exact bonds:
10-11 11-12 12-23 15-16 19-26
normalized bonds:
1-2 1-6 2-3 3-4 4-5 5-6 16-17 16-21 17-18 18-19 19-20 20-21 23-24 23-25

Connectivity :

12:3 E exact RC ring/chain

Match level:

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS

#### => FILE HCAPLUS

FILE 'HCAPLUS' ENTERED AT 18:31:46 ON 20 NOV 2009
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 20 Nov 2009 VOL 151 ISS 22

FILE LAST UPDATED: 19 Nov 2009 (20091119/ED)

REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2009

USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2009

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2009.

CAS Information Use Policies apply and are available at:

## http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.

During November, try the new LSUS format of legal status information in the  ${\rm CA/CAplus}$  family databases for free! Complete details on the

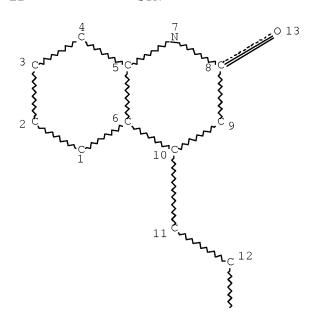
#### Page 4 of 49

number of free displays and other databases participating in this offer appear in NEWS 10.

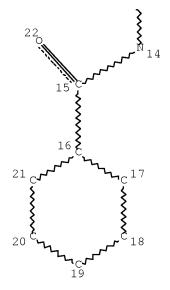
'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

## => D STAT QUE L30

L2 STR



Page 1-A



Page 2	2-A
--------	-----

NODE ATTRIBUTES:

NSPEC	IS	R	AT	1
NSPEC	IS	R	AT	2
NSPEC	IS	R	AT	3
NSPEC	TS	R	ΑТ	4

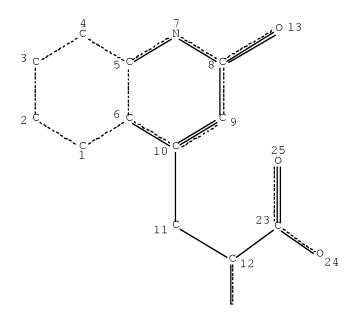
```
AT 5
NSPEC
       IS R
       IS R
                 ΑT
NSPEC
       IS R
                 ΑT
                      7
NSPEC
NSPEC
       IS R
                 ΑT
                      8
NSPEC
       IS R
                 ΑT
                     9
NSPEC
       IS R
                 AT 10
NSPEC
       IS C
                 AT 11
NSPEC
       IS C
                 AT
                    12
NSPEC
       IS C
                 AT 13
NSPEC
       IS C
                    14
                 AT
       IS C
NSPEC
                 ΑT
                    15
NSPEC
       IS R
                 ΑT
                    16
NSPEC
       IS R
                 ΑT
                    17
NSPEC
      IS R
                 AT 18
NSPEC
      IS R
                 AT 19
NSPEC
       IS R
                 ΑT
                    20
NSPEC
       IS R
                 ΑT
                    21
       IS C
NSPEC
                 ΑT
DEFAULT MLEVEL IS ATOM
MLEVEL IS CLASS AT 11 12 13 14 15 22
DEFAULT ECLEVEL IS LIMITED
```

#### GRAPH ATTRIBUTES:

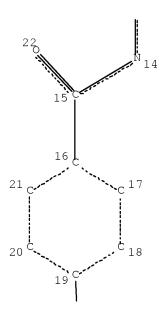
RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 22

#### STEREO ATTRIBUTES: NONE

L4 99 SEA FILE=REGISTRY SSS FUL L2 L5 STR



Page 1-A



Page 2-A



Page 3-	-A			
NODE AT	TRIE	BUTES:		
NSPEC	IS	R	ΑT	1
NSPEC	IS	R	ΑT	2
NSPEC	IS	R	ΑT	3
NSPEC	IS	R	ΑT	4
NSPEC	IS	R	ΑT	5
NSPEC	IS	R	ΑT	6
NSPEC	IS	R	ΑT	7
NSPEC	IS	R	ΑT	8
NSPEC	IS	R	ΑT	9
NSPEC	IS	R	ΑT	10
NSPEC	IS	С	ΑT	11
NSPEC	IS	С	ΑT	12
NSPEC	IS	С	ΑT	13
NSPEC	IS	С	ΑT	14
NSPEC	IS	С	ΑT	15
NSPEC	IS	R	ΑT	16
NSPEC	IS	R	ΑT	17
NSPEC	IS	R	ΑT	18
NSPEC	IS	R	ΑT	19
NSPEC	IS	R	ΑT	20
NSPEC	IS	R	ΑT	21
NSPEC	IS	С	ΑT	22
NSPEC	IS	С	ΑT	23
NSPEC	IS	С	ΑT	24
NSPEC	IS	С	ΑT	25
NSPEC	IS	С	ΑT	26

Page 7 of 49

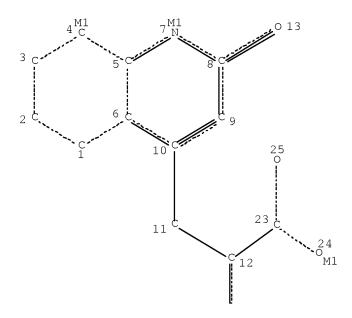
DEFAULT MLEVEL IS ATOM
MLEVEL IS CLASS AT 11 12 13 14 15 22 23 24 25 26
DEFAULT ECLEVEL IS LIMITED

#### GRAPH ATTRIBUTES:

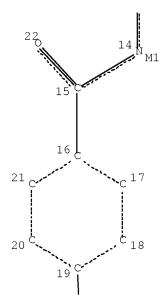
RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 26

## STEREO ATTRIBUTES: NONE

L7 60 SEA FILE=REGISTRY SUB=L4 SSS FUL L5 L13 STR



Page 1-A



```
Page 2-A
```

```
Cl 26
```

```
Page 3-A
  NODE ATTRIBUTES:
NODE ATTRIBUTES:

HCOUNT IS M1 AT 4

HCOUNT IS M1 AT 14

HCOUNT IS M1 AT 14

HCOUNT IS M1 AT 24

NSPEC IS R AT 1

NSPEC IS R AT 3

NSPEC IS R AT 4

NSPEC IS R AT 5

NSPEC IS R AT 5

NSPEC IS R AT 6

NSPEC IS R AT 7

NSPEC IS R AT 7

NSPEC IS R AT 8

NSPEC IS R AT 9

NSPEC IS R AT 10

NSPEC IS R AT 11

NSPEC IS R AT 10

NSPEC IS C AT 11

NSPEC IS C AT 11

NSPEC IS C AT 12

NSPEC IS R AT 16

NSPEC IS C AT 15

NSPEC IS R AT 16

NSPEC IS R AT 17

NSPEC IS R AT 16

NSPEC IS R AT 17

NSPEC IS R AT 17

NSPEC IS R AT 17

NSPEC IS R AT 19

NSPEC IS R AT 20

NSPEC IS R AT 21

NSPEC IS R AT 21

NSPEC IS R AT 22

NSPEC IS C AT 22

NSPEC IS C AT 23

NSPEC IS C AT 25

NSPEC IS C AT 25
  HCOUNT IS M1 AT
  NSPEC IS C AT 26
  CONNECT IS E3 RC AT 12
   DEFAULT MLEVEL IS ATOM
  MLEVEL IS CLASS AT 11 12 13 14 15 22 23 24 25 26
  DEFAULT ECLEVEL IS LIMITED
```

#### GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 26

STEREO ATTRIBUTES: NONE

STEREO	ATTRIBUTE	15: I	NONE				
L14	35	SEA	FILE=REGISTRY	SUB=L7	SSS FUL	L13	
L15	356	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L14
L16	57021	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	SALIVA+PFT/CT OR
		SAL	IVA?/BI				
L17	11	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L16(L)ACCELERATION/OBI
L18	1	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L15 AND L17
L19	630	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	XEROSTOMIA+OLD, PFT/CT
L20	2	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L15 AND L19

Page 9 of 49

L23	112	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	NAGAMOTO H?/AU
L24	78	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	KOHASHI M?/AU
L25	1964	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	OKA H?/AU
L26	3	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	L15 AND L16
L27	3	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	L18 OR L20 OR L26
L28	3	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	L27 AND ((L23 OR L24
		OR L25))			
L29	1	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	L23 AND L24 AND L25
L30	3	SEA FILE=HCAPLUS	SPE=ON ABB=	ON PLU=ON	L28 OR L29

#### => FILE MEDLINE

FILE 'MEDLINE' ENTERED AT 18:32:00 ON 20 NOV 2009

FILE LAST UPDATED: 18 Nov 2009 (20091118/UP). FILE COVERS 1949 TO DATE.

MEDLINE and LMEDLINE have been updated with the 2009 Medical Subject Headings (MeSH) vocabulary and tree numbers from the U.S. National Library of Medicine (NLM). Additional information is available at

http://www.nlm.nih.gov/pubs/techbull/nd08/nd08\_medline\_data\_changes\_2009.html.

On February 21, 2009, MEDLINE was reloaded. See HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

See HELP RANGE before carrying out any RANGE search.

## => D STAT QUE L50

L21	1	SEA	FILE=REGISTR	Y SPE=ON	ABB=ON	PLU=ON	"4-QUINOLINEPROPANOIC
		AC:	ID, $A-((4-CHL)$	OROBENZO	YL)AMINO	)-1,2-DI	HYDRO-2-OXO-"/CN
L31		SEL	PLU=ON L21	1- NAME	:	4 TERMS	
L32	228	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L31
L33	228	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L21 OR L32
L34	26444	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	SALIVA/CT
L37	3041	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	XEROSTOMIA/CT
L40	1	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L33 AND L37
L41	1	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L33 AND L34
L42	1	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L40 OR L41
L43	8594	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	"SJOGREN'S SYNDROME"/C
		Τ					
L44	3	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L33 AND L43
L45	3	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L42 OR L44
L46	34	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	NAGAMOTO H?/AU
L47	19	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	KOHASHI M?/AU
L48	932	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	OKA H?/AU
L50	2	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L45 AND ((L46 OR L47
OR L48))							

#### => FILE WPIX BIOSIS

FILE 'WPIX' ENTERED AT 18:32:54 ON 20 NOV 2009 COPYRIGHT (C) 2009 THOMSON REUTERS

FILE 'BIOSIS' ENTERED AT 18:32:54 ON 20 NOV 2009 Copyright (c) 2009 The Thomson Corporation

=> D STAT QUE L67

L64 82 SEA NAGAMOTO H?/AU

L65 73 SEA KOHASHI M?/AU L66 3368 SEA OKA H?/AU

L67 1 SEA L64 AND L65 AND L66

#### => FILE EMBASE

FILE 'EMBASE' ENTERED AT 18:33:02 ON 20 NOV 2009 Copyright (c) 2009 Elsevier B.V. All rights reserved.

FILE COVERS 1974 TO 20 Nov 2009 (20091120/ED)

EMBASE was reloaded on March 30, 2008.

EMBASE is now updated daily. SDI frequency remains weekly (default) and biweekly.

This file contains CAS Registry Numbers for easy and accurate substance identification.

Beginning January 2008, Elsevier will no longer provide EMTREE codes as part of the EMTREE thesaurus in EMBASE. Please update your current-awareness alerts (SDIs) if they contain EMTREE codes.

For further assistance, please contact your local helpdesk.

=> D STAT	QUE L	85					
L21	1	SEA	FILE=REGIST	RY SPE=O	N ABB=0	N PLU=O	N "4-QUINOLINEPROPANOIC
		AC:	ID, $A-((4-CH))$	LOROBENZ	OYL)AMIN	(0) - 1, 2 - D	OIHYDRO-2-OXO-"/CN
L68		SEL	PLU=ON L23	1 1- NAM	E :	4 TERM	S
L69	381	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L68
L70	381	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L21 OR L69
L71	17167	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	XEROSTOMIA/CT
L72	9448	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	SJOEGREN SYNDROME/CT
L74	7	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L70 AND L71
L75	12	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L70 AND L72
L77	2	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L75 AND MANAGEMENT/TI
L78	5	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L75 NOT (EYE OR
		RHE	JMATOLOGIC)/	ΓΙ			
L79	7	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L77 OR L78
L80	9	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L74 OR L79
L81	35	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	NAGAMOTO H?/AU
L82	15	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	KOHASHI M?/AU
L83	863	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	OKA H?/AU
L85	2	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L80 AND (L81 OR L82 OR
		L83	)				

#### => DUP REMOVE L30 L50 L67 L85

FILE 'HCAPLUS' ENTERED AT 18:33:28 ON 20 NOV 2009
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'MEDLINE' ENTERED AT 18:33:28 ON 20 NOV 2009

FILE 'WPIX' ENTERED AT 18:33:28 ON 20 NOV 2009 COPYRIGHT (C) 2009 THOMSON REUTERS

FILE 'EMBASE' ENTERED AT 18:33:28 ON 20 NOV 2009

Copyright (c) 2009 Elsevier B.V. All rights reserved.

PROCESSING COMPLETED FOR L50 PROCESSING COMPLETED FOR L67 PROCESSING COMPLETED FOR L85

L86 3 DUP REMOVE L30 L50 L67 L85 (5 DUPLICATES REMOVED)

ANSWERS '1-3' FROM FILE HCAPLUS

=> D L86 IBIB ABS HITSTR 1-3

L86 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2009 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2009:392963 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 151:395986

TITLE: Efficacy and safety of rebamipide for the treatment of

dry mouth symptoms in patients with Sjoegren's
syndrome: a double-blind placebo-controlled

multicenter trial

AUTHOR(S): Sugai, Susumu; Takahashi, Hiroki; Ohta, Shuji;

Nishinarita, Makoto; Takei, Masami; Sawada, Shigemasa;

Yamaji, Ken; Oka, Hiroshi; Umehara,

Hisanori; Koni, Ichiro; Sugiyama, Eiji; Nishiyama,

Susumu; Kawakami, Atsushi

CORPORATE SOURCE: Kanazawa Medical University, Ishikawa, Japan

SOURCE: Modern Rheumatology (2009), 19(2), 114-124

CODEN: MROHA4; ISSN: 1439-7595

PUBLISHER: Springer Japan

DOCUMENT TYPE: Journal LANGUAGE: English

The effects of rebamipide on dry mouth and salivary secretion in Sjoegren's syndrome patients were investigated in a double-blind placebo-controlled study. Rebamipide (100 mg TID) or placebo was administered for eight weeks and patient-assessed improvement of dry mouth and increase in salivary secretion measured by the Saxon test were evaluated. At two, four, and eight weeks, dry mouth improvement rates were, resp., 26.0%, 44.0%, and 46.9% for rebamipide and 20.0%, 27.1%, and 39.1% for placebo, and mean increases in salivary secretion were, resp., 0.14, 0.24, and 0.35 q for rebamipide and 0.03, 0.09, and 0.17 q for placebo, indicating higher values in the rebamipide group for both parameters at all timepoints but no significant differences between the two groups. Anal. by baseline characteristics suggested a statistically significant salivary secretion increasing effect of rebamipide in cases of primary Sjoegren's syndrome. No difference in the incidence of adverse events was seen between the two groups, confirming the safety of rebamipide. As a salivary secretion increasing effect was strongly suggested in cases of primary Sjoegren's syndrome, further study on the administration of rebamipide for the treatment of dry mouth in patients with Sjoegren's syndrome is required.

IT 90098-04-7, Mucosta

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Mucosta was effective in increasing salivary secretion but

had no effect on dry mouth symptoms in patient with Sjogren's syndrome)

RN 90098-04-7 HCAPLUS

CN 4-Quinoline propanoic acid,  $\alpha$ -[(4-chlorobenzoyl)amino]-1,2-dihydro-2-oxo- (CA INDEX NAME)

REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L86 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2009 ACS on STN DUPLICATE 2

ACCESSION NUMBER: 2008:292244 HCAPLUS Full-text

DOCUMENT NUMBER: 148:576370

TITLE: Effective treatment with oral administration of rebamipide in a mouse model of Sjogren's syndrome

AUTHOR(S): Kohashi, Masayuki; Ishimaru, Naozumi;

Arakaki, Rieko; Hayashi, Yoshio

CORPORATE SOURCE: University of Tokushima Graduate School, Tokushima,

Japan

SOURCE: Arthritis & Rheumatism (2008), 58(2), 389-400

CODEN: ARHEAW; ISSN: 0004-3591

PUBLISHER: John Wiley & Sons, Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Objective: To determine whether oral administration of rebamipide, a mucosal protective agent, is effective in the treatment of Sjogren's syndrome (SS) in the NFS/sld mouse model of the disease. Methods: NFS/sld mice were given daily oral doses of rebamipide (0.3 mg/kg of body weight or 3 mg/kg) or vehicle alone starting from the age of 4 wk to the age of 8 wk. The volume of saliva and tears was monitored during and after treatment. After the final dose, histol. features of the tissues, TUNEL + apoptotic duct cells in affected glands, T cell and cytokine function, and levels of Ig isotypes and serum autoantibodies were examined Results: The 3-mg/kg dose of rebamipide prevented the development of autoimmune lesions. average volume of saliva in rebamipide-treated mice was significantly higher than that in control mice. We found decreased TUNEL + apoptotic duct cells in the salivary and lacrimal glands of rebamipide-treated mice as compared with control mice. Rebamipide treatment suppressed the activation of CD4 + T cells and Th1 cytokines (interleukin-2, interferon- $\gamma$ ) associated with impaired NF- $\kappa$ B activity. Production of serum autoantibodies, IgM, and IgG1 was clearly inhibited. Conclusion: Our findings demonstrate the efficacy of oral administration of rebamipide in the treatment of SS. Rebamipide represents a new therapeutic strategy for the treatment of patients with sicca symptoms caused by SS, as well as for patients with other diseases.

IT 90098-04-7, Rebamipide

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(effective treatment with oral administration of rebamipide in mouse model of Sjogren's syndrome)

RN 90098-04-7 HCAPLUS

CN

4-Quinolinepropanoic acid,  $\alpha$ -[(4-chlorobenzoyl)amino]-1,2-dihydro-2-oxo- (CA INDEX NAME)

OS.CITING REF COUNT: 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD

(3 CITINGS)

REFERENCE COUNT: 48 THERE ARE 48 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L86 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2009 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 2005:120780 HCAPLUS Full-text

DOCUMENT NUMBER: 142:183519

TITLE: Carbostyril derivatives for accelerating

salivation

INVENTOR(S): Nagamoto, Hisashi; Kohashi, Masayuki

; Oka, Hiroshi

PATENT ASSIGNEE(S): Otsuka Pharmaceutical Co., Ltd., Japan; St. Marianna

University School of Medicine

SOURCE: PCT Int. Appl., 36 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005011811	A1	20050210	WO 2004-JP9992	20040707
W: AE, AG,	AL, AM, AT	r, AU, AZ, BA,	BB, BG, BR, BW,	BY, BZ, CA, CH,
CN, CO,	CR, CU, CZ	Z, DE, DK, DM,	DZ, EC, EE, EG,	ES, FI, GB, GD,
GE, GH,	GM, HR, HU	J, ID, IL, IN,	IS, JP, KE, KG,	KP, KR, KZ, LC,
LK, LR,	LS, LT, LU	J, LV, MA, MD,	MG, MK, MN, MW,	MX, MZ, NA, NI,
NO, NZ,	OM, PG, PH	H, PL, PT, RO,	RU, SC, SD, SE,	SG, SK, SL, SY,
TJ, TM,	TN, TR, TT	r, TZ, UA, UG,	US, UZ, VC, VN,	YU, ZA, ZM, ZW
RW: BW, GH,	GM, KE, LS	S, MW, MZ, NA,	SD, SL, SZ, TZ,	UG, ZM, ZW, AM,
AZ, BY,	KG, KZ, MD	O, RU, TJ, TM,	AT, BE, BG, CH,	CY, CZ, DE, DK,
EE, ES,	FI, FR, GB	B, GR, HU, IE,	IT, LU, MC, NL,	PL, PT, RO, SE,
SI, SK,	TR, BF, BJ	J, CF, CG, CI,	CM, GA, GN, GQ,	GW, ML, MR, NE,
SN, TD,	TG			
EP 1648563	A1	20060426	EP 2004-747458	20040707
EP 1648563	B1	20070919		
R: AT, BE,	CH, DE, DK	K, ES, FR, GB,	GR, IT, LI, LU,	NL, SE, MC, PT,
IE, SI,	FI, RO, CY	, TR, BG, CZ,	EE, HU, PL, SK	

CN 1859948	A	20061108	CN	2004-80027975	5	20040707
JP 2006528	662 T	20061221	JP	2006-521775		20040707
AT 373502	T	20071015	AT	2004-747458		20040707
KR 2006037	408 A	20060503	KR	2006-701933		20060127
US 2007011:	2026 A1	20070517	US	2006-566214		20060127
PRIORITY APPLN.	INFO.:		JP	2003-282691	A	20030730
			JP	2004-21808	A	20040129
			WO	2004-JP9992	W	20040707

OTHER SOURCE(S): MARPAT 142:183519

AB An oral pharmaceutical composition for accelerating salivation and prophylaxis and/or treatment of xerostomia or hyposalivation comprises as an active ingredient a carbostyril compound or a pharmaceutically acceptable salt thereof. For example, a mixture containing 2-(4-chlorobenzoylamino)-3-(2-quinolon-4-yl)propionic acid (Rebamipide) 150 g, Avicel 40 g, corn starch 30 g, and magnesium stearate 2 g was tableted and film coated with a composition containing hydroxypropyl Me cellulose 10 g, polyethylene glycol 6000 3 g, castor oil 40 g, and methanol 40 g. Tablets containing 100 mg Rebamipide per tablet were orally administered three times per day immediately after a meal to patients having Sjogren's syndrome. An increase of salivation was observed with the effectiveness of 52.4% after 4 wk and 61.9% after 8 wk of administration.

IT 90098-04-7, Rebamipide

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(oral carbostyril derivs. for accelerating salivation)

RN 90098-04-7 HCAPLUS

CN 4-Quinoline propanoic acid,  $\alpha$ -[(4-chlorobenzoyl)amino]-1,2-dihydro-2-oxo- (CA INDEX NAME)

OS.CITING REF COUNT: 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## Serial#: 10/566,214 STRUCTURE SEARCH

#### => FILE HCAPLUS

FILE 'HCAPLUS' ENTERED AT 18:33:53 ON 20 NOV 2009
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 20 Nov 2009 VOL 151 ISS 22
FILE LAST UPDATED: 19 Nov 2009 (20091119/ED)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2009
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2009

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2009.

CAS Information Use Policies apply and are available at:

## http://www.cas.org/legal/infopolicy.html

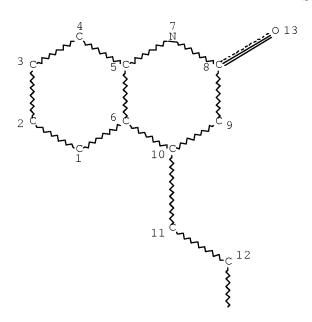
This file contains CAS Registry Numbers for easy and accurate substance identification.

During November, try the new LSUS format of legal status information in the  ${\rm CA/CAplus}$  family databases for free! Complete details on the number of free displays and other databases participating in this offer appear in NEWS 10.

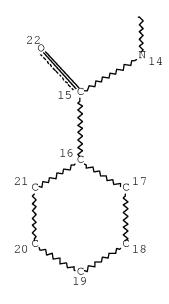
'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

## => D STAT QUE L53

L2 STR



Page 1-A



Page	2-A				
NODE .	ATTRII	3U:	res:		
NSPEC	IS	R		ΑT	1
NSPEC	IS	R		ΑT	2
NSPEC	IS	R		ΑT	3
NSPEC	IS	R		ΑT	4
NSPEC	IS	R		ΑT	5
NSPEC	IS	R		ΑT	6
NSPEC	IS	R		ΑT	7
NSPEC	IS	R		ΑT	8
NSPEC	IS	R		ΑT	9
NSPEC	IS	R		ΑT	10

Page 17 of 49

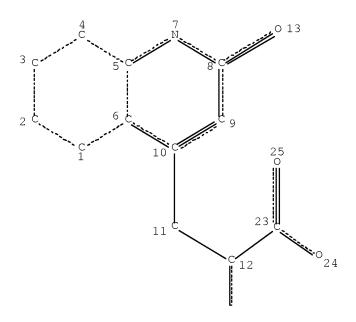
IS C AT 11 NSPEC NSPEC IS C AT 12 NSPEC IS C ΑT 13 NSPEC IS C ΑT 14 NSPEC IS C ΑT 15 16 NSPEC IS R ΑT NSPEC IS R ΑT 17 NSPEC IS R ΑT 18 NSPEC IS R ΑT 19 IS R 20 NSPEC ATNSPEC IS R ΑT 21 NSPEC IS C ΑT DEFAULT MLEVEL IS ATOM MLEVEL IS CLASS AT 11 12 13 14 15 22 DEFAULT ECLEVEL IS LIMITED

## GRAPH ATTRIBUTES:

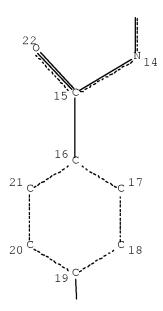
RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 22

#### STEREO ATTRIBUTES: NONE

L4 99 SEA FILE=REGISTRY SSS FUL L2 L5 STR



Page 1-A



Page 2-A



Page 3-	-A			
NODE AT	TRIE	BUTES:		
NSPEC	IS	R	AT	1
NSPEC	IS	R	AT	2
NSPEC	IS	R	ΑT	3
NSPEC	IS	R	ΑT	4
NSPEC	IS	R	ΑT	5
NSPEC	IS	R	ΑT	6
NSPEC	IS	R	ΑT	7
NSPEC	IS	R	ΑT	8
NSPEC	IS	R	ΑT	9
NSPEC	IS	R	ΑT	10
NSPEC	IS	С	ΑT	11
NSPEC	IS	С	ΑT	12
NSPEC	IS	С	ΑT	13
NSPEC	IS	С	ΑT	14
NSPEC	IS	С	ΑT	15
NSPEC	IS	R	ΑT	16
NSPEC	IS	R	ΑT	17
NSPEC	IS	R	ΑT	18
NSPEC	IS	R	ΑT	19
NSPEC	IS	R	ΑT	20
NSPEC	IS	R	ΑT	21
NSPEC	IS	С	ΑT	22
NSPEC	IS	С	ΑT	23
NSPEC	IS	С	ΑT	24
NSPEC	IS	С	ΑT	25
NSPEC	IS	С	ΑT	26

Page 19 of 49

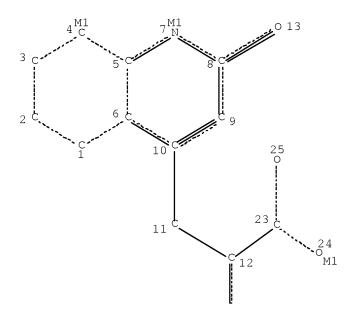
DEFAULT MLEVEL IS ATOM
MLEVEL IS CLASS AT 11 12 13 14 15 22 23 24 25 26
DEFAULT ECLEVEL IS LIMITED

#### GRAPH ATTRIBUTES:

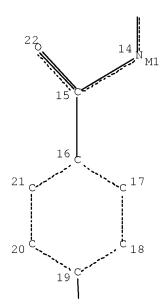
RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 26

## STEREO ATTRIBUTES: NONE

L7 60 SEA FILE=REGISTRY SUB=L4 SSS FUL L5 L13 STR



Page 1-A



```
Page 2-A
```

d<sub>1 26</sub>

```
Page 3-A
  NODE ATTRIBUTES:
NODE ATTRIBUTES:

HCOUNT IS M1 AT 4

HCOUNT IS M1 AT 14

HCOUNT IS M1 AT 14

HCOUNT IS M1 AT 24

NSPEC IS R AT 1

NSPEC IS R AT 3

NSPEC IS R AT 4

NSPEC IS R AT 5

NSPEC IS R AT 5

NSPEC IS R AT 6

NSPEC IS R AT 7

NSPEC IS R AT 7

NSPEC IS R AT 8

NSPEC IS R AT 9

NSPEC IS R AT 10

NSPEC IS R AT 11

NSPEC IS R AT 10

NSPEC IS C AT 11

NSPEC IS C AT 11

NSPEC IS C AT 12

NSPEC IS R AT 16

NSPEC IS C AT 15

NSPEC IS R AT 16

NSPEC IS R AT 17

NSPEC IS R AT 16

NSPEC IS R AT 17

NSPEC IS R AT 17

NSPEC IS R AT 17

NSPEC IS R AT 19

NSPEC IS R AT 20

NSPEC IS R AT 21

NSPEC IS R AT 21

NSPEC IS R AT 22

NSPEC IS C AT 22

NSPEC IS C AT 23

NSPEC IS C AT 25

NSPEC IS C AT 25
  HCOUNT IS M1 AT 4
                                                                    AT 26
  NSPEC IS C
  CONNECT IS E3 RC AT 12
   DEFAULT MLEVEL IS ATOM
  MLEVEL IS CLASS AT 11 12 13 14 15 22 23 24 25 26
  DEFAULT ECLEVEL IS LIMITED
```

## GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 26

STEREO ATTRIBUTES: NONE

STEREO	ATTRIBUTE	4S: I	NONE				
L14	35	SEA	FILE=REGISTRY	SUB=L7	SSS FUL	L13	
L15	356	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L14
L16	57021	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	SALIVA+PFT/CT OR
		SAL	IVA?/BI				
L17	11	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L16(L)ACCELERATION/OBI
L18	1	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L15 AND L17
L19	630	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	XEROSTOMIA+OLD, PFT/CT
L20	2	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L15 AND L19

L26	3	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L15 AND L16
L27	3	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L18 OR L20 OR L26
L51	3703	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	SJOGREN SYNDROME+OLD, P
		FT/C	CT				
L52	4	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L15 AND L51
L53	4	SEA	FILE=HCAPLUS	SPE=ON	ABB=ON	PLU=ON	L27 OR L52

#### => FILE MEDLINE

FILE 'MEDLINE' ENTERED AT 18:34:11 ON 20 NOV 2009

FILE LAST UPDATED: 18 Nov 2009 (20091118/UP). FILE COVERS 1949 TO DATE.

MEDLINE and LMEDLINE have been updated with the 2009 Medical Subject Headings (MeSH) vocabulary and tree numbers from the U.S. National Library of Medicine (NLM). Additional information is available at

http://www.nlm.nih.gov/pubs/techbull/nd08/nd08\_medline\_data\_changes\_2009.html.

On February 21, 2009, MEDLINE was reloaded. See HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

See HELP RANGE before carrying out any RANGE search.

#### => FILE HCAPLUS

FILE 'HCAPLUS' ENTERED AT 18:34:19 ON 20 NOV 2009
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 20 Nov 2009 VOL 151 ISS 22

FILE LAST UPDATED: 19 Nov 2009 (20091119/ED)

REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2009

USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2009

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2009.

CAS Information Use Policies apply and are available at:

http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.

During November, try the new LSUS format of legal status information in the CA/CAplus family databases for free! Complete details on the number of free displays and other databases participating in this

#### Page 22 of 49

offer appear in NEWS 10. 'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

#### => S L53 NOT L30

L87 1 L53 NOT L30

#### => FILE MEDLINE

FILE 'MEDLINE' ENTERED AT 18:34:42 ON 20 NOV 2009

FILE LAST UPDATED: 18 Nov 2009 (20091118/UP). FILE COVERS 1949 TO DATE.

MEDLINE and LMEDLINE have been updated with the 2009 Medical Subject Headings (MeSH) vocabulary and tree numbers from the U.S. National Library of Medicine (NLM). Additional information is available at

http://www.nlm.nih.gov/pubs/techbull/nd08/nd08\_medline\_data\_changes\_2009.html.

On February 21, 2009, MEDLINE was reloaded. See HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

See HELP RANGE before carrying out any RANGE search.

#### => D STAT QUE L45

L21	1	SEA	FILE=REGISTRY	Y SPE=ON	ABB=ON	PLU=ON	"4-QUINOLINEPROPANOIC					
	ACID, A-((4-CHLOROBENZOYL)AMINO)-1,2-DIHYDRO-2-OXO-"/CN											
L31		SEL	PLU=ON L21	1- NAME	:	4 TERMS						
L32	228	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L31					
L33	228	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L21 OR L32					
L34	26444	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	SALIVA/CT					
L37	3041	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	XEROSTOMIA/CT					
L40	1	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L33 AND L37					
L41	1	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L33 AND L34					
L42	1	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L40 OR L41					
L43	8594	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	"SJOGREN'S SYNDROME"/C					
		Τ										
L44	3	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L33 AND L43					
<b>L45</b>	3	SEA	FILE=MEDLINE	SPE=ON	ABB=ON	PLU=ON	L42 OR L44					

#### => S L45 NOT L50

L88 1 L45 NOT L50

#### => FILE WPIX BIOSIS

FILE 'WPIX' ENTERED AT 18:35:23 ON 20 NOV 2009 COPYRIGHT (C) 2009 THOMSON REUTERS

FILE 'BIOSIS' ENTERED AT 18:35:23 ON 20 NOV 2009 Copyright (c) 2009 The Thomson Corporation

#### => D STAT QUE L60

, , ,	20	00					
L21	1	SEA	FILE=REGISTRY	SPE=ON	ABB=ON	PLU=ON	"4-QUINOLINEPROPANOIC
		AC:	ID, $A-(4-CHLO)$	ROBENZOY	L)AMINO)	-1,2-DIH	YDRO-2-OXO-"/CN
L54		SEL	PLU=ON L21	1- NAME	:	4 TERMS	
L55	379	SEA	L54				
L56	379	SEA	L21 OR L55				
L57	3824	SEA	SALIVA? (10A) (2	ACCEL? O	R INCREA	.S?)	
1.60	0	SEA	I.56 (30A) I.57				

#### => D STAT QUE L62

L21	1	SEA FILE=REGISTRY SPE=ON ABB=ON PLU=ON "4-QUINOLINEPROPANOIC
		ACID, A-((4-CHLOROBENZOYL)AMINO)-1,2-DIHYDRO-2-OXO-"/CN
L54		SEL PLU=ON L21 1- NAME : 4 TERMS
L55	379	SEA L54
L56	379	SEA L21 OR L55
L61	4470	SEA XEROSTO? OR DRY(3A) MOUTH?
L62	0	SEA L56 AND L61

#### => FILE EMBASE

FILE 'EMBASE' ENTERED AT 18:35:44 ON 20 NOV 2009 Copyright (c) 2009 Elsevier B.V. All rights reserved.

FILE COVERS 1974 TO 20 Nov 2009 (20091120/ED)

EMBASE was reloaded on March 30, 2008.

EMBASE is now updated daily. SDI frequency remains weekly (default) and biweekly.

This file contains CAS Registry Numbers for easy and accurate substance identification.

Beginning January 2008, Elsevier will no longer provide EMTREE codes as part of the EMTREE thesaurus in EMBASE. Please update your current-awareness alerts (SDIs) if they contain EMTREE codes.

For further assistance, please contact your local helpdesk.

#### => D STAT QUE L80

L21	1	SEA	FILE=REGIST	RY SPE=O	N ABB=O	N PLU=O	N "4-QUINOLINEPROPANOIC
		AC	ID, A-((4-CH	LOROBENZ	OYL) AMIN	O)-1,2-D	IHYDRO-2-OXO-"/CN
L68		SEL	PLU=ON L21	l 1- NAM	E :	4 TERM	S
L69	381	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L68
L70	381	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L21 OR L69
L71	17167	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	XEROSTOMIA/CT
L72	9448	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	SJOEGREN SYNDROME/CT
L74	7	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L70 AND L71
L75	12	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L70 AND L72
L77	2	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L75 AND MANAGEMENT/TI
L78	5	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L75 NOT (EYE OR
		RHEU	JMATOLOGIC)/I	ΓI			
L79	7	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L77 OR L78
<b>T80</b>	9	SEA	FILE=EMBASE	SPE=ON	ABB=ON	PLU=ON	L74 OR L79

## => S L80 NOT L85

L89 7 L80 NOT L85

#### => DUP REMOVE L87 L88 L89

FILE 'HCAPLUS' ENTERED AT 18:36:18 ON 20 NOV 2009
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'MEDLINE' ENTERED AT 18:36:18 ON 20 NOV 2009

FILE 'EMBASE' ENTERED AT 18:36:18 ON 20 NOV 2009

Copyright (c) 2009 Elsevier B.V. All rights reserved.

PROCESSING COMPLETED FOR L87 PROCESSING COMPLETED FOR L88 PROCESSING COMPLETED FOR L89

8 DUP REMOVE L87 L88 L89 (1 DUPLICATE REMOVED) L90

> ANSWER '1' FROM FILE HCAPLUS ANSWER '2' FROM FILE MEDLINE ANSWERS '3-8' FROM FILE EMBASE

#### L90 ANSWER 1 OF 8 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2009:1267017 HCAPLUS Full-text

DOCUMENT NUMBER:

151:478529 Topical LFA-1 antagonists for use in localized TITLE:

treatment of immune related disorders

Burnier, John; Gadek, Thomas INVENTOR(S):

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 85pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.					KIND DATE				APPLICATION NO.						DATE			
US 20090258070					A1 20091015			US 2009-386359						20090415				
WO	WO 2009128934			A1 20091022					WO 2009-US2389					20090415				
	W:	ΑE,	AG,	AL,	AM,	AO,	ΑT,	ΑU,	AZ,	BA,	BB,	BG,	BH,	BR,	BW,	BY,	BZ,	
		CA,	CH,	CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DO,	DZ,	EC,	EE,	EG,	ES,	
		FI,	GB,	GD,	GE,	GH,	GM,	GT,	HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	
		KG,	KM,	KN,	KP,	KR,	KΖ,	LA,	LC,	LK,	LR,	LS,	LT,	LU,	LY,	MA,	MD,	
		ME,	MG,	MK,	MN,	MW,	MX,	MY,	MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	
		PL,	PT,	RO,	RS,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	ST,	SV,	SY,	ТJ,	
		TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	ZA,	ZM,	ZW			
	RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HR,	HU,	
		ΙE,	IS,	ΙT,	LT,	LU,	LV,	MC,	MK,	MT,	NL,	NO,	PL,	PT,	RO,	SE,	SI,	
		SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	$\mathrm{ML}_{m{\prime}}$	MR,	ΝE,	SN,	
		TD,	TG,	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	
		ZW,	AM,	ΑZ,	BY,	KG,	KΖ,	MD,	RU,	ΤJ,	TM							
TTTC	7 7 7 7	T INT	TNEO							TTC O	$\cap \cap \circ$	1521	ΛD		D 2	0000	115	

PRIORITY APPLN. INFO.:

US 2008-45240P

This invention provides specifically formulated LFA-1 antagonists or pharmaceutically acceptable salts thereof that are suitable for topical delivery. In particular, the LFA-1 antagonists are particularly well suited for localized treatment by having a rapid systemic clearance rate. The invention also encompasses methods of treatment and prevention of immune related disorders using the LFA-1 topical formulations of the present invention.

INCL 424489000; 514314000; 514311000; 514300000; 514448000; 514009000

63-6 (Pharmaceuticals)

Section cross-reference(s): 1

ΙT Alopecia

> Angiogenesis inhibitors Anti-inflammatory agents

Antiarthritics Antiasthmatics

Antimicrobial agents

Antioxidants

Antirheumatic agents

Antiulcer agents

Antiviral agents

```
Arthralgia
Asthma
Beeswax
Blood plasma
Chelating agents
Chronic obstructive pulmonary disease
Complexing agents
Crohn disease
Eczema
Edema
Emulsions
Graves' disease
Human
Immune disease
Immunomodulators
Lubricants
Oral drug delivery systems
Permeation enhancers
Pharmaceutical creams
Pharmaceutical emulsions
Pharmaceutical foams
Pharmaceutical gels
Pharmaceutical ointments
Pharmaceutical pastes
Pharmaceutical powders
Pharmaceutical solutions
Pharmaceutical sprays
Pharmaceutical suspensions
Psoriasis
Pulmonary fibrosis
Retinal disease
Retinitis
Rheumatoid arthritis
  Sjogren syndrome
Surfactants
T cell
Topical drug delivery systems
Ulcerative colitis
Uveitis
Volatile substances
Wetting agents
   (topical LFA-1 antagonists for use in localized treatment of immune
   related disorders)
59865-13-3, Cyclosporine 59985-21-6, Diquafosol 90098-04-7,
Rebamipide 245466-70-0 851784-48-0 851784-50-4 851784-54-8
851784-56-0 851784-64-0 851784-72-0 915397-46-5 915397-52-3 915397-63-6 915397-90-9 915397-99-8 1025967-78-5 1026240-04-9
1148048-06-9 1148048-11-6 1148048-14-9 1148048-17-2
                                                              1148048-24-1
1148048-27-4 1148048-29-6 1148048-36-5 1148048-38-7
RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU
(Therapeutic use); BIOL (Biological study); USES (Uses)
   (topical LFA-1 antagonists for use in localized treatment of immune
   related disorders)
90098-04-7, Rebamipide
RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU
(Therapeutic use); BIOL (Biological study); USES (Uses)
   (topical LFA-1 antagonists for use in localized treatment of immune
   related disorders)
90098-04-7 HCAPLUS
4-Quinolinepropanoic acid, \alpha-[(4-chlorobenzoyl)amino]-1,2-dihydro-2-
```

ΤТ

ΤТ

RN

CN

L90 ANSWER 2 OF 8 MEDLINE on STN DUPLICATE 1

ACCESSION NUMBER: 2009123510 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 18841440

TITLE: Sjogren's syndrome: promising, new treatment options

besides nizatidine.

AUTHOR: Kapoor Shailendra

SOURCE: Modern rheumatology / the Japan Rheumatism Association,

(2009) Vol. 19, No. 1, pp. 100-1. Electronic Publication:

2008-10-08.

Journal code: 100959226. ISSN: 1439-7595.

COMMENT: Comment on: Mod Rheumatol. 2008;18(5):455-9. PubMed ID:

18478182

PUB. COUNTRY: Japan
DOCUMENT TYPE: Commentary

Letter

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200906

ENTRY DATE: Entered STN: 10 Feb 2009

Last Updated on STN: 2 Jul 2009 Entered Medline: 30 Jun 2009

CONTROLLED TERM: \*Alanine: AA, analogs & derivatives

Alanine: TU, therapeutic use
Antioxidants: TU, therapeutic use

\*Flax

Histamine H2 Antagonists: TU, therapeutic use

Humans

\*Nizatidine: TU, therapeutic use

\*Plant Preparations: TU, therapeutic use

\*Quinolones: TU, therapeutic use

\*Seeds

\*Sjogren's Syndrome: DT, drug therapy

Treatment Outcome

CAS REGISTRY NO.: 111911-87-6 (rebamipide); 56-41-7 (Alanine);

76963-41-2 (Nizatidine)

CHEMICAL NAME: 0 (Antioxidants); 0 (Histamine H2 Antagonists); 0 (Plant

Preparations); 0 (Quinolones)

L90 ANSWER 3 OF 8 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights reserved on STN ACCESSION NUMBER: 2008439484 EMBASE Full-text Rebamipide as an important adjunctive tool in the TITLE: management of rheumatologic disorders: Comment on the article by Kohashi et al. AUTHOR: Kapoor, Shailendra, Dr. (correspondence) CORPORATE SOURCE: University of Illinois at Chicago. Arthritis and Rheumatism, (September 2008) Vol. 58, No. 9, SOURCE: pp. 2923. Refs: 9 ISSN: 0004-3591 CODEN: ARHEAW PUBLISHER: John Wiley and Sons Inc., P.O.Box 18667, Newark, NJ 07191-8667, United States. United States COUNTRY: DOCUMENT TYPE: Journal; Letter 031 Arthritis and Rheumatism FILE SEGMENT: 037 Drug Literature Index LANGUAGE: English ENTRY DATE: Entered STN: 17 Oct 2008 Last Updated on STN: 17 Oct 2008 CONTROLLED TERM: Medical Descriptors: angiogenesis aphthous ulcer: DT, drug therapy Behcet disease digestive system ulcer: DT, drug therapy digestive system ulcer: PC, prevention drug efficacy human letter priority journal \*rheumatic disease: DT, drug therapy Sjoegren syndrome: DT, drug therapy ulcerative colitis: DT, drug therapy xerostomia: DT, drug therapy CONTROLLED TERM: Drug Descriptors: misoprostol: CM, drug comparison nonsteroid antiinflammatory agent \*rebamipide: CM, drug comparison \*rebamipide: DT, drug therapy \*rebamipide: PD, pharmacology CAS REGISTRY NO.: (misoprostol) 59122-46-2, 59122-48-4; (rebamipide ) 111911-87-6 L90 ANSWER 4 OF 8 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights reserved on STN ACCESSION NUMBER: 2008502560 EMBASE Full-text TITLE: Treatment of Dry Eye Disease by the Non-Ophthalmologist. AUTHOR: Foulks, Gary N., Dr. (correspondence) Department of Ophthalmology and Visual Sciences, University CORPORATE SOURCE: of Louisville School of Medicine, 301 East Muhammad Ali Boulevard, Louisville, KY 40202, United States. gnfoul01@lo uisville.edu SOURCE: Rheumatic Disease Clinics of North America, (November 2008)

Vol. 34, No. 4, pp. 987-1000.

Refs: 72

ISSN: 0889-857X CODEN: RDCAEK

PUBLISHER: W.B. Saunders, Independence Square West, Philadelphia, PA

19106-3399, United States.

PUBLISHER IDENT.: S 0889-857X(08)00079-3

COUNTRY: United States

DOCUMENT TYPE: Journal; General Review; (Review)

FILE SEGMENT: 012 Ophthalmology

031 Arthritis and Rheumatism 037 Drug Literature Index 038 Adverse Reactions Titles

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 3 Dec 2008

Last Updated on STN: 3 Dec 2008

ABSTRACT: Physicians caring for patients who have Sjogren's syndrome often face a particularly difficult task in managing the dry eye that occurs with this disease. The discomfort produced by the condition and the fluctuation of vision attendant to tear film instability are often the most annoying of the clinical symptoms. The understanding of dry eye disease in both its clinical expression and underlying etiopathology has expanded over the past 10 years with implications for management and therapy. The array of potential treatments both topical and systemic has evolved to provide a much more targeted and effective arsenal from which the clinician may choose. .COPYRGT. 2008 Elsevier Inc. All rights reserved.

CONTROLLED TERM: Medical Descriptors:

antiinflammatory activity
cataract: SI, side effect

clinical trial

disease classification

disease severity
drug efficacy

\*dry eye: DT, drug therapy \*dry eye: ET, etiology \*dry eye: SU, surgery \*dry eye: TH, therapy

eye inflammation: DT, drug therapy

glaucoma: SI, side effect

hormonal therapy

human

keratoconjunctivitis sicca: DT, drug therapy

lacrimation pathogenesis priority journal

review

rosacea: DT, drug therapy

sebaceous gland disease: DT, drug therapy

serum

Sjoegren syndrome

superinfection: SI, side effect

treatment outcome

zerostomia: DT, drug therapy

CONTROLLED TERM: Drug Descriptors:

12 sulfodehydroabietic acid: CT, clinical trial 12 sulfodehydroabietic acid: DT, drug therapy 12 sulfodehydroabietic acid: TP, topical drug

administration

acetylcysteine: DT, drug therapy

acetylcysteine: TP, topical drug administration

antiinflammatory agent: DT, drug therapy

artificial tear: DT, drug therapy

```
artificial tear: TP, topical drug administration
                    cevimeline: DT, drug therapy
                    cevimeline: PO, oral drug administration
                    corticosteroid: CB, drug combination
                    corticosteroid: DT, drug therapy
                    corticosteroid: TP, topical drug administration
                    cyclosporin A: AE, adverse drug reaction
                    cyclosporin A: CB, drug combination
                    cyclosporin A: DT, drug therapy
                    cyclosporin A: PD, pharmacology
                    cyclosporin A: TP, topical drug administration
                    diquafosol: CT, clinical trial
                    diquafosol: DT, drug therapy
                    diquafosol: TP, topical drug administration
                    estratest: DT, drug therapy
                    estrogen: DT, drug therapy
                    estrogen: TP, topical drug administration
                    eye drops: DT, drug therapy
                    eye drops: TP, topical drug administration
                    idestrin estradiol
                    immunomodulating agent: DT, drug therapy
                    immunomodulating agent: TP, topical drug administration
                    lipid emulsion: CT, clinical trial
                    lipid emulsion: DT, drug therapy
                    lipid emulsion: TP, topical drug administration
                    loteprednol etabonate: DT, drug therapy
                    loteprednol etabonate: TP, topical drug administration
                    lubricating agent: CB, drug combination
                    lubricating agent: DT, drug therapy
                    lubricating agent: TP, topical drug administration
                    omega 3 fatty acid: CB, drug combination
                    omega 3 fatty acid: DT, drug therapy
                    omega 3 fatty acid: PO, oral drug administration
                    omega 3 fatty acid: TP, topical drug administration
                    optive
                    pilocarpine: CT, clinical trial
                    pilocarpine: DT, drug therapy
                    pilocarpine: PO, oral drug administration
                     rebamipide: CT, clinical trial
                      rebamipide: DT, drug therapy
                      rebamipide: TP, topical drug administration
                    refresh endura
                    soothe
                    steroid: AE, adverse drug reaction
                    steroid: DT, drug therapy
                    steroid: TP, topical drug administration
                    systane
                    testosterone: CT, clinical trial
                    testosterone: DT, drug therapy
                    testosterone: TP, topical drug administration
                    tetracycline: DT, drug therapy
                    theratears nutrition
                    unclassified drug
SUPPLEMENTARY TERM: Cyclosporine; Dry eye; Secretagogues; Sjogren's syndrome
                   (12 sulfodehydroabietic acid) 33159-27-2, 86408-72-2;
                    (acetylcysteine) 616-91-1; (cevimeline) 107220-27-9,
                    107220-28-0, 107233-08-9, 153504-70-2; (cyclosporin A)
                    59865-13-3, 63798-73-2; (diquafosol) 211427-08-6;
                    (loteprednol etabonate) 82034-46-6; (pilocarpine) 148-72-1,
                    54-71-7, 92-13-7; (rebamipide)
                    111911-87-6; (testosterone) 58-22-0; (tetracycline)
```

CAS REGISTRY NO.:

23843-90-5, 60-54-8, 64-75-5

CHEMICAL NAME: (1) estratest; (2) evoxac; (3) idestrin estradiol; (4)

lotemax; (5) optive; (6) refresh endura; (7) restasis; (8)

salagen; (9) soothe; (10) systane; (11) theratears

nutrition

COMPANY NAME: (1) Solvay (United States); (2) Daiichi Seiyaku (United

States); (3) nascent (United States); (4) Bausch and Lomb (United States); (5) Allergan (United States); (6) Allergan

(United States); (7) Allergan (United States); (8) MGI (United States); (9) Bausch and Lomb (United States); (10) Alcon (United States); (11) advanced research (United

States); Inspire (United States); Novartis (United States);

Otsuka (United States)

L90 ANSWER 5 OF 8 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights

reserved on STN

ACCESSION NUMBER: 2008312638 EMBASE Full-text

TITLE: Current and prospective treatment options for Sjogren, s

syndrome.

AUTHOR: Sugai, Susumu, Dr. Prof. (correspondence)

CORPORATE SOURCE: Kudo General Hospital, Kanazawa Medical University,

Kanazawa, Japan. sussugai@helen.ocn.ne.jp

AUTHOR: Masaki, Yasafumi

CORPORATE SOURCE: Department of Hematology and Immunology, Kanazawa Medical

University, 1-1 Daigaku, Uchinada, Kahoku-gun, Ishikawa

920-0293, Japan. yasum@kanazawa-med.ac.jp

AUTHOR: Sugai, Susumu, Dr. Prof. (correspondence)

CORPORATE SOURCE: Kudo General Hospital, Kanazawa University, Kanazawa, Japan

. sussugai@helen.ocn.ne.jp

SOURCE: Expert Review of Clinical Immunology, (Jul 2008) Vol. 4,

No. 4, pp. 469-479.

Refs: 87

ISSN: 1744-666X

COUNTRY: United Kingdom

DOCUMENT TYPE: Journal; General Review; (Review)

FILE SEGMENT: 011 Otorhinolaryngology

012 Ophthalmology

026 Immunology, Serology and Transplantation 030 Clinical and Experimental Pharmacology

037 Drug Literature Index 038 Adverse Reactions Titles

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 10 Jul 2008

Last Updated on STN: 10 Jul 2008

treatment of Sjogren's syndrome. .COPYRGT. 2008 Expert Reviews Ltd.

ABSTRACT: Sjogren's syndrome is both an organ-specific and a systemic autoimmune disease. Patients with dry eyes and dry mouth usually receive symptomatic treatment. Topical cyclosporine A may be administered to patients with dry eyes. The efficacy of muscarinic agonists, such as pilocarpine and cevimeline, in increasing salivary and lacrimal flow and improving patient symptoms has been well documented. Mild immunomodulatory or immunosuppressive medication is preferable to prevent the mild progressive course of the disorders. Other patients who show aggressive progression to organ disorders should be managed differently. High doses of corticosteroids, immunosuppressive agents or new biological agents, such as antibodies to CD20, are required for these patients. The elimination of B lymphocytes by anti-CD20 antibodies may be highly beneficial; however, the development of more selective drugs based on the pathogenesis of the disease is required for the future

CONTROLLED TERM: Medical Descriptors:

```
abdominal pain: SI, side effect
                    B lymphocyte
                    clinical trial
                    disease course
                    drug dose comparison
                    drug dose increase
                    drug efficacy
                    drug induced headache: SI, side effect
                    drug mechanism
                    drug megadose
                    drug safety
                    drug selectivity
                    drug targeting
                    drug tolerability
                    dry eye: DT, drug therapy
                    dry eye: SU, surgery
                    human
                    immunopathogenesis
                    keratoconjunctivitis sicca: DT, drug therapy
                    keratoconjunctivitis sicca: SU, surgery
                    low drug dose
                    monotherapy
                    mucosa associated lymphoid tissue lymphoma: DT, drug
                    therapy
                    nausea: SI, side effect
                    nonhuman
                    plasmapheresis
                    recommended drug dose
                    regulatory T lymphocyte
                    review
                    salivation
                    side effect: SI, side effect
                      *Sjoegren syndrome: DT, drug therapy
                      *Sjoegren syndrome: ET, etiology
                    sweating
                    tear flow
                    unspecified side effect: SI, side effect
                    vasculitis: DT, drug therapy
                    vasculitis: TH, therapy
                    viral gene delivery system
                    viral gene therapy
                     zerostomia: DT, drug therapy
CONTROLLED TERM:
                    Drug Descriptors:
                    alpha interferon: CT, clinical trial
                    alpha interferon: DO, drug dose
                    alpha interferon: DT, drug therapy
                    alpha interferon: PO, oral drug administration
                    alpha interferon: PD, pharmacology
                    antirheumatic agent: DT, drug therapy
                    aquaporin 1: DT, drug therapy
                    artificial tear: DT, drug therapy
                    CD20 antibody: DT, drug therapy
                    CD40 ligand monoclonal antibody: DT, drug therapy
                    CD40 ligand monoclonal antibody: IP, intraperitoneal drug
                    administration
                    CD40 ligand monoclonal antibody: PD, pharmacology
                    cevimeline: AE, adverse drug reaction
                    cevimeline: CT, clinical trial
                    cevimeline: DO, drug dose
                    cevimeline: DT, drug therapy
                    cevimeline: PO, oral drug administration
```

```
cevimeline: PD, pharmacology
                    corticosteroid: CB, drug combination
                    corticosteroid: DO, drug dose
                    corticosteroid: DT, drug therapy
                    corticosteroid: PO, oral drug administration
                    cyclophosphamide: CB, drug combination
                    cyclophosphamide: DT, drug therapy
                    cyclophosphamide: IV, intravenous drug administration
                    cyclosporin A: DT, drug therapy
                    cyclosporin A: TP, topical drug administration
                    doxorubicin: CB, drug combination
                    doxorubicin: DT, drug therapy
                    etanercept: CT, clinical trial
                    etanercept: DT, drug therapy
                    gefarnate: DT, drug therapy
                    gefarnate: PD, pharmacology
                    hydroxychloroquine: DT, drug therapy
                    hydroxychloroquine: PD, pharmacology
                    immunomodulating agent: DT, drug therapy
                    immunosuppressive agent: DT, drug therapy
                    infliximab: CT, clinical trial
                    infliximab: DT, drug therapy
                    interleukin 10: DT, drug therapy
                    interleukin 10: PD, pharmacology
                    mizoribine: AE, adverse drug reaction
                    mizoribine: CT, clinical trial
                    mizoribine: DT, drug therapy
                    mizoribine: PD, pharmacology
                    muscarinic agent: DT, drug therapy
                    parvovirus vector
                    pilocarpine: AE, adverse drug reaction
                    pilocarpine: CT, clinical trial
                    pilocarpine: DO, drug dose
                    pilocarpine: DT, drug therapy
                    pilocarpine: PO, oral drug administration
                    pilocarpine: PD, pharmacology
                    pilocarpine: SC, subcutaneous drug administration
                    placebo
                    prednisolone: DT, drug therapy
                    prednisone: CB, drug combination
                    prednisone: DT, drug therapy
                      rebamipide: AE, adverse drug reaction
                     rebamipide: CT, clinical trial
                      rebamipide: DT, drug therapy
                      rebamipide: PO, oral drug administration
                      rebamipide: PD, pharmacology
                    rituximab: CT, clinical trial
                    rituximab: CB, drug combination
                    rituximab: DO, drug dose
                    rituximab: DT, drug therapy
                    sni 2011
                    tumor necrosis factor alpha inhibitor: DT, drug therapy
                    unindexed drug
                    vincristine: CB, drug combination
                    vincristine: DT, drug therapy
CAS REGISTRY NO.:
                    (aquaporin 1) 146410-94-8, 149348-86-7; (cevimeline)
                    107220-27-9, 107220-28-0, 107233-08-9, 153504-70-2;
                    (cyclophosphamide) 50-18-0; (cyclosporin A) 59865-13-3,
                    63798-73-2; (doxorubicin) 23214-92-8, 25316-40-9;
                    (etanercept) 185243-69-0, 200013-86-1; (gefarnate) 51-77-4;
                    (hydroxychloroquine) 118-42-3, 525-31-5; (infliximab)
```

170277-31-3; (mizoribine) 50924-49-7; (pilocarpine) 148-72-1, 54-71-7, 92-13-7; (prednisolone) 50-24-8;

(prednisone) 53-03-2; (rebamipide)

111911-87-6; (rituximab) 174722-31-7; (vincristine)

57-22-7

CHEMICAL NAME: sni 2011

L90 ANSWER 6 OF 8 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights

reserved on STN

ACCESSION NUMBER: 2008131921 EMBASE Full-text

TITLE: Primary Sjogren's Syndrome: Current and Prospective

Therapies.

AUTHOR: Thanou-Stavraki, Aikaterini; James, Judith A., Dr.

(correspondence)

CORPORATE SOURCE: Arthritis and Immunology Program, Oklahoma Medical Research

Foundation, Oklahoma City, OK, United States. jamesj@omrf.o

uhsc.edu

Thanou-Stavraki, Aikaterini; James, Judith A., Dr. **AUTHOR:** 

(correspondence)

Department of Medicine, University of Oklahoma Health CORPORATE SOURCE:

Sciences Center, Oklahoma City, OK, United States.

jamesj@omrf.ouhsc.edu

AUTHOR: James, Judith A., Dr. (correspondence)

Department of Pathology, University of Oklahoma Health CORPORATE SOURCE:

Sciences Center, Oklahoma City, OK, United States.

jamesj@omrf.ouhsc.edu

Seminars in Arthritis and Rheumatism, (Apr 2008) Vol. 37, SOURCE:

No. 5, pp. 273-292.

Refs: 251

ISSN: 0049-0172 CODEN: SAHRBF

S 0049-0172(07)00113-8 PUBLISHER IDENT.:

COUNTRY: United States DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 031 Arthritis and Rheumatism

> 037 Drug Literature Index 038 Adverse Reactions Titles

Internal Medicine 006

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 7 Apr 2008

Last Updated on STN: 7 Apr 2008

Objective: To summarize data on existing and experimental therapies for ABSTRACT: primary Sjogren's syndrome (pSS), referring both to sicca syndrome and to other systemic disease manifestations. Methods: Relevant English and non-English articles acquired through Medline were reviewed. Results: pSS usually has a benign clinical course, centered on sicca features and general musculoskeletal manifestations, and is managed symptomatically. However, a subset of patients develops more severe extraglandular disease that warrants close monitoring and aggressive treatment. For dry eyes and mouth, nonpharmacologic measures to preserve secretions, and tear and saliva substitutes, offer some symptomatic relief. Muscarinic agonists and topical cyclosporine yield well-documented improvement in ocular sicca features. Although traditional antirheumatic drugs are used empirically for polyarthritis and other Sjogren's symptoms, their efficacy in pSS overall and as disease-modifying agents is limited. For the potential severe, nonexocrine manifestations complicating pSS, standard high-dose immunosuppression is used. Among the biologic agents already examined in pSS, those targeting tumor necrosis factor (TNF)- $\alpha$  failed to demonstrate significant benefit. Nonetheless, rituximab and other B-cell-depleting therapies appear promising. Conclusions: Treatment of pSS patients with severe extraglandular disease

should differ from that of patients with predominantly sicca features and/or general muscoloskeletal manifestations. pSS treatment is mainly symptomatic, primarily directed against sicca complaints. The traditional anti-rheumatic agents show limited efficacy in the systemic process and use of systemic  ${\tt TNF-}\alpha$  inhibitors has been very disappointing. B cell depleting treatments and other newer biologic therapies appear more promising. .COPYRGT. 2008 Elsevier Inc. All rights reserved.

```
CONTROLLED TERM:
                    Medical Descriptors:
                    antiinflammatory activity
                    article
                    atopic dermatitis: DT, drug therapy
                    B cell lymphoma
                    clinical feature
                    clinical trial
                    disease severity
                    drug efficacy
                    drug indication
                    drug safety
                    dry eye: DT, drug therapy
                    dry eye: SI, side effect
                    eye discharge: DT, drug therapy
                    glomerulonephritis
                    hematologic disease
                    human
                    immunosuppressive treatment
                    keratoconjunctivitis sicca: SU, surgery
                    lacrimal gland disease: DT, drug therapy
                    lacrimation
                    low drug dose
                    lymphocyte depletion
                    MEDLINE
                    menopausal syndrome: DT, drug therapy
                    neurologic disease
                    patient monitoring
                    polyarthritis: DT, drug therapy
                    priority journal
                    salivation
                      *Sjoegren syndrome: DT, drug therapy
                    topical treatment
                    treatment outcome
                    vasculitis
                    viral gene delivery system
                      xerostomia: DT, drug therapy
                      xerostomia: TH, therapy
CONTROLLED TERM:
                    Drug Descriptors:
                    12 sulfodehydroabietic acid: DT, drug therapy
                    alpha interferon: DO, drug dose
                    alpha interferon: DT, drug therapy
                    alpha interferon: PO, oral drug administration
                    alpha interferon: PA, parenteral drug administration
                    alpha interferon: PD, pharmacology
                    antihistaminic agent: AE, adverse drug reaction
                    antiinflammatory agent: DT, drug therapy
                    antirheumatic agent: DT, drug therapy
                    beta adrenergic receptor blocking agent: AE, adverse drug
                    reaction
                    cevimeline: CM, drug comparison
                    cevimeline: DT, drug therapy
                    corticosteroid: DT, drug therapy
```

corticosteroid: TP, topical drug administration

```
cyclosporin A: CT, clinical trial
                    cyclosporin A: CR, drug concentration
                    cyclosporin A: DT, drug therapy
                    cyclosporin A: PK, pharmacokinetics
                    cyclosporin A: PD, pharmacology
                    cyclosporin A: TP, topical drug administration
                    diquafosol: DT, drug therapy
                    diquafosol: PD, pharmacology
                    diquafosol: TP, topical drug administration
                    disease modifying antirheumatic drug: DT, drug therapy
                    diuretic agent: AE, adverse drug reaction
                    estratest: DT, drug therapy
                    eye drops: DT, drug therapy
                    eye drops: TP, topical drug administration
                    gefarnate: DT, drug therapy
                    immunosuppressive agent: DT, drug therapy
                    muscarinic agent: DT, drug therapy
                    nonsteroid antiinflammatory agent: DT, drug therapy
                    nonsteroid antiinflammatory agent: TP, topical drug
                    administration
                    pilocarpine: CM, drug comparison
                    pilocarpine: DT, drug therapy
                    pimecrolimus: DT, drug therapy
                    pimecrolimus: TP, topical drug administration
                      rebamipide: DT, drug therapy
                    recombinant interleukin 10: DT, drug therapy
                    rituximab: DT, drug therapy
                    tacrolimus: DT, drug therapy
                    tacrolimus: TP, topical drug administration
                    testosterone: DT, drug therapy
                    testosterone: TP, topical drug administration
                    testosterone: TD, transdermal drug administration
                    tricyclic antidepressant agent: AE, adverse drug reaction
                    tumor necrosis factor alpha: EC, endogenous compound
                    tumor necrosis factor alpha receptor: DT, drug therapy
                    unindexed drug
                    vasoactive intestinal polypeptide: DT, drug therapy
                    vasoactive intestinal polypeptide: PD, pharmacology
                    (12 sulfodehydroabietic acid) 33159-27-2, 86408-72-2;
CAS REGISTRY NO.:
                    (cevimeline) 107220-27-9, 107220-28-0, 107233-08-9,
                    153504-70-2; (cyclosporin A) 59865-13-3, 63798-73-2;
                    (diquafosol) 211427-08-6; (gefarnate) 51-77-4;
                    (pilocarpine) 148-72-1, 54-71-7, 92-13-7; (pimecrolimus)
                    137071-32-0; (rebamipide) 111911-87-6;
                    (rituximab) 174722-31-7; (tacrolimus) 104987-11-3;
                    (testosterone) 58-22-0; (vasoactive intestinal polypeptide)
                    37221-79-7
CHEMICAL NAME:
                    (1) evoxac; (2) restasis; (3) salagen; evoxac
COMPANY NAME:
                    (1) Daiichi Seiyaku (United States); (2) Allergan (United
                    States); (3) Novartis (United States)
L90 ANSWER 7 OF 8 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights
     reserved on STN
ACCESSION NUMBER: 2008073013 EMBASE
                                          Full-text
TITLE:
                    Pharmacological management of dry eye in the elderly
                    patient.
                   Foulks, Gary N., Dr. Prof. (correspondence)
AUTHOR:
CORPORATE SOURCE: Department of Ophthalmology and Vision Science, University
                    of Louisville School of Medicine, Louisville, KY, United
                    States.
```

AUTHOR: Foulks, Gary N., Dr. Prof. (correspondence)

CORPORATE SOURCE: 301 E. Muhammad Ali Boulevard, Louisville, KY 40202, United

States.

SOURCE: Drugs and Aging, (2008) Vol. 25, No. 2, pp. 105-118.

Refs: 98

ISSN: 1170-229X CODEN: DRAGE6

COUNTRY: New Zealand

DOCUMENT TYPE: Journal; General Review; (Review)

FILE SEGMENT: 012 Ophthalmology

017 Public Health, Social Medicine and Epidemiology

O20 Gerontology and Geriatrics O37 Drug Literature Index

038 Adverse Reactions Titles

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 6 Mar 2008

Last Updated on STN: 6 Mar 2008

ABSTRACT: Dry eye disease is a common and increasingly prevalent condition particularly associated with advancing age and postmenopausal women.

Epidemiological studies identify prevalence rates ranging from 7% in the US to 33% in the Asian population. Research increasingly identifies risk factors of increasing age, female sex, smoking, use of video display terminals and use of certain medications as well as environmental stresses as aggravating factors for the disease. Basic and clinical investigations provide cumulative evidence of hyperosmolarity of the tear film and ocular surface/lacrimal gland inflammation as pathogenic features of dry eye disease. A decline in systemic and local levels of sex hormones is associated with advancing age and advancing disease. Pharmacological therapeutic interventions include enhanced lubricants and anti-inflammatory drugs such as topical corticosteroids and ciclosporin (cyclosporine A). Secretagogues and hormonal supplementation are potential future therapies. The increased understanding of the contributing and pathogenetic factors responsible for dry eye provides a rationale for multiple therapeutic options for this multi-factorial disease. In the elderly patient it is important to recognize the physical and cognitive limitations that will influence the selection of appropriate topical medication. .COPYRGT. 2008 Adis Data Information BV. All rights reserved.

CONTROLLED TERM: Medical Descriptors:

Asian

cataract: SI, side effect

clinical trial cognition

\*dry eye: DT, drug therapy \*dry eye: EP, epidemiology

elderly care
emulsion

environmental exposure

fluorescence Hispanic

hormone substitution

human

inflammation

intraocular pressure

lubrication

multifactorial genetic disorder

nonhuman osmolarity prevalence

priority journal

review
risk factor

```
sex difference
                    side effect: SI, side effect
                    social behavior
                    staining
                    tear film
                      xerostomia: DT, drug therapy
CONTROLLED TERM:
                    Drug Descriptors:
                    12 sulfodehydroabietic acid: CT, clinical trial
                    artificial tear: IT, drug interaction
                    artificial tear: DT, drug therapy
                    cevimeline: CT, clinical trial
                    cevimeline: CM, drug comparison
                    cevimeline: DT, drug therapy
                    corticosteroid: AE, adverse drug reaction
                    corticosteroid: DT, drug therapy
                    corticosteroid: PD, pharmacology
                    corticosteroid: TP, topical drug administration
                    cyclosporin A: CT, clinical trial
                    cyclosporin A: IT, drug interaction
                    cyclosporin A: DT, drug therapy
                    cyclosporin A: TP, topical drug administration
                    diquafosol: CT, clinical trial
                    diquafosol: DT, drug therapy
                    diquafosol: PD, pharmacology
                    diquafosol: TP, topical drug administration
                    duramycin: CT, clinical trial
                    estratest: CT, clinical trial
                    estratest: DT, drug therapy
                    estratest: TP, topical drug administration
                    freshkote
                    loteprednol etabonate: CT, clinical trial
                    loteprednol etabonate: DT, drug therapy
                    omega 3 fatty acid: DT, drug therapy
                    pilocarpine: CM, drug comparison
                    pilocarpine: DT, drug therapy
                      rebamipide: CT, clinical trial
                      rebamipide: DT, drug therapy
                      rebamipide: TP, topical drug administration
                    refresh endura
                    restoryl
                    soothe
                    systane
                    (12 sulfodehydroabietic acid) 33159-27-2, 86408-72-2;
CAS REGISTRY NO.:
                    (cevimeline) 107220-27-9, 107220-28-0, 107233-08-9,
                    153504-70-2; (cyclosporin A) 59865-13-3, 63798-73-2;
                    (diquafosol) 211427-08-6; (duramycin) 1391-36-2;
                    (loteprednol etabonate) 82034-46-6; (pilocarpine) 148-72-1,
                    54-71-7, 92-13-7; (rebamipide)
                    111911-87-6
CHEMICAL NAME:
                    (1) estratest; (2) evoxac; (3) freshkote; (4) moli 1901;
                    (5) refresh endura; (6) restasis; (7) restoryl; (8)
                    salagen; (9) soothe; (10) systane
COMPANY NAME:
                    (1) Solvay (United States); (2) Daiichi Seiyaku (United
                    States); (3) Focus (United States); (4) lantibio (United
                    States); (5) Allergen (United States); (6) Allergen (United
                    States); (7) Bausch and Lomb (United States); (8) MGI
                    (United States); (9) Bausch and Lomb (United States); (10)
                    Alcon (United States); Inspire (United States); ISTA
                    (United States); Otsuka (United States)
```

reserved on STN

ACCESSION NUMBER: 2008312479 EMBASE Full-text

TITLE: Management and therapy of dry eye disease: Report

of the management and therapy subcommittee of the

international Dry Eye WorkShop (2007).

AUTHOR: Pflugfelder, Stephen C., Dr. (correspondence)

CORPORATE SOURCE: Ophthalmology-Ocular Surf Ctr., Cullen Eye Institute, 6565

Fannin Street NC 205, Houston, TX 77030, United States.

stevenp@bcm.tmc.edu

AUTHOR: Lemp, Michael A.

CORPORATE SOURCE: 4000 Cathedral Avenue NW, Washington, DC 20016, United

States. malemp@lempdc.com

AUTHOR: Geerling, Gerd; Kinoshita, Shigero; McCulley, James;

Nelson, Daniel; Novack, Gary N.; Shimazaki, Jun; Wilson,

Clive

SOURCE: Ocular Surface, (Apr 2007) Vol. 5, No. 2, pp. 163-178.

Refs: 185

ISSN: 1542-0124

COUNTRY: United States

DOCUMENT TYPE: Journal; Conference Article; (Conference paper)

FILE SEGMENT: 012 Ophthalmology

030 Clinical and Experimental Pharmacology

037 Drug Literature Index038 Adverse Reactions Titles

039 Pharmacy052 Toxicology

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 10 Jul 2008

Last Updated on STN: 10 Jul 2008

ABSTRACT: The members of the Management and Therapy Subcommittee assessed current dry eye therapies. Each member wrote a succinct evidence-based review on an assigned aspect of the topic, and the final report was written after review by and with consensus of all subcommittee members and the entire Dry Eye WorkShop membership. In addition to Its own review of the literature, the Subcommittee reviewed the Dry Eye Preferred Practice Patterns of the American Academy of Ophthalmology and the International Task Force (ITF) Delphi Panel on Dry Eye. The Subcommittee favored the approach taken by the ITF, whose recommended treatments were based on level of disease severity. The recommendations of the Subcommittee are based on a modification of the ITF severity grading scheme, and suggested treatments were chosen from a menu of therapies for which evidence of therapeutic effect had been presented. .COPYRGT. 2007 Ethis Communications, Inc.

CONTROLLED TERM: Medical Descriptors:

absence of side effects: SI, side effect

autotransplantation

blepharitis: DT, drug therapy burning sensation: SI, side effect

clinical trial conference paper contact lens dose response

drug dose comparison drug dose increase drug efficacy drug formulation drug mechanism

drug megadose
drug safety

drug tolerability

arug toler

```
drug withdrawal
*dry eye: DT, drug therapy
*dry eye: SU, surgery
eye disease: SI, side effect
human
hyperhidrosis: SI, side effect
keratoconjunctivitis sicca: DT, drug therapy
low drug dose
monotherapy
nonhuman
osmolarity
rheumatoid arthritis: DT, drug therapy
rosacea: DT, drug therapy
salivary gland
Schirmer test
  Sjoegren syndrome: DT, drug therapy
systemic disease: SI, side effect
tear film
unspecified side effect: SI, side effect
Drug Descriptors:
12 sulfodehydroabietic acid: CT, clinical trial
12 sulfodehydroabietic acid: DT, drug therapy
12 sulfodehydroabietic acid: TP, topical drug
administration
15 hydroxyicosatetraenoic acid: DT, drug therapy
15 hydroxyicosatetraenoic acid: PD, pharmacology
15 hydroxyicosatetraenoic acid: TP, topical drug
administration
androgen: CT, clinical trial
androgen: DT, drug therapy
androgen: TP, topical drug administration
angiogenesis inhibitor: DT, drug therapy
antiinfective agent: DT, drug therapy
antiinflammatory agent: DT, drug therapy
artificial tear: CT, clinical trial
artificial tear: CB, drug combination
artificial tear: DT, drug therapy
benzalkonium chloride: TO, drug toxicity
cevimeline: AE, adverse drug reaction
cevimeline: CT, clinical trial
cevimeline: DO, drug dose
cevimeline: DT, drug therapy
cevimeline: PO, oral drug administration
corticosteroid: CT, clinical trial
corticosteroid: DT, drug therapy
cyclosporin A: AE, adverse drug reaction
cyclosporin A: CT, clinical trial
cyclosporin A: DO, drug dose
cyclosporin A: DT, drug therapy
cyclosporin A: TP, topical drug administration
de 089
diquafosol: CT, clinical trial
diquafosol: DT, drug therapy
diquafosol: TP, topical drug administration
doxycycline: CT, clinical trial
doxycycline: DO, drug dose
doxycycline: DT, drug therapy
dwelle
edetic acid: TO, drug toxicity
endura
essential fatty acid: CT, clinical trial
```

CONTROLLED TERM:

```
essential fatty acid: DT, drug therapy
                    fluorometholone: CT, clinical trial
                    fluorometholone: CB, drug combination
                    fluorometholone: DT, drug therapy
                    flurbiprofen: CT, clinical trial
                    flurbiprofen: CB, drug combination
                    flurbiprofen: DT, drug therapy
                    gefarnate: DT, drug therapy
                    gefarnate: TP, topical drug administration
                    genteal
                    hypotears: CT, clinical trial
                    hypotears: DT, drug therapy
                    hypotears: PR, pharmaceutics
                    hypotears pf
                    loteprednol etabonate: CT, clinical trial
                    loteprednol etabonate: DT, drug therapy
                    methylprednisolone: AE, adverse drug reaction
                    methylprednisolone: CT, clinical trial
                    methylprednisolone: DT, drug therapy
                    methylprednisolone: TP, topical drug administration
                    minocycline: CT, clinical trial
                    minocycline: DT, drug therapy
                    optive
                    pilocarpine: AE, adverse drug reaction
                    pilocarpine: CT, clinical trial
                    pilocarpine: DT, drug therapy
                    pilocarpine: PO, oral drug administration
                    placebo
                      rebamipide: CT, clinical trial
                      rebamipide: DT, drug therapy
                      rebamipide: PD, pharmacology
                      rebamipide: TP, topical drug administration
                    refresh
                    refresh free
                    soothe
                    systane
                    tacrolimus: CT, clinical trial
                    tacrolimus: DT, drug therapy
                    tears natural free
                    testosterone: CT, clinical trial
                    testosterone: DT, drug therapy
                    testosterone: TP, topical drug administration
                    tetracycline derivative: CT, clinical trial
                    tetracycline derivative: DT, drug therapy
                    tetracycline derivative: PD, pharmacology
                    tetryzoline: DT, drug therapy
                    theratears
                    unindexed drug
                    visine pure tears
                    (12 sulfodehydroabietic acid) 33159-27-2, 86408-72-2; (15
CAS REGISTRY NO.:
                    hydroxyicosatetraenoic acid) 73180-00-4; (benzalkonium
                    chloride) 66331-30-4, 78244-97-0, 81181-32-0; (cevimeline)
                    107220-27-9, 107220-28-0, 107233-08-9, 153504-70-2;
                    (cyclosporin A) 59865-13-3, 63798-73-2; (diquafosol)
                    211427-08-6; (doxycycline) 10592-13-9, 17086-28-1,
                    564-25-0; (edetic acid) 150-43-6, 60-00-4; (essential fatty
                    acid) 11006-87-4; (fluorometholone) 426-13-1;
                    (flurbiprofen) 5104-49-4; (gefarnate) 51-77-4; (loteprednol
                    etabonate) 82034-46-6; (methylprednisolone) 6923-42-8,
                    83-43-2; (minocycline) 10118-90-8, 11006-27-2, 13614-98-7;
                    (pilocarpine) 148-72-1, 54-71-7, 92-13-7; (
```

rebamipide) 111911-87-6; (tacrolimus) 104987-11-3; (testosterone) 58-22-0; (tetryzoline) 522-48-5, 84-22-0

CHEMICAL NAME:

(1) de 089; (2) de 089; (3) dwelle; (4) endura; (5) genteal; (6) hypotears pf; (7) ins 365; (8) ins 365; (9) lotemax; (10) opc 12759; (11) opc 12759 ; (12) optive; (13) refresh free; (14) refresh; (15)

soothe; (16) systame; (17) tears natural free; (18)

theratears; (19) visine pure tears

COMPANY NAME:

(1) Inspire (United States); (2) Santen (Japan); (3) Dry Eye (United States); (4) Allergan (United States); (5) Novartis (United States); (6) Novartis (United States); (7) Inspire (United States); (8) Santen (Japan); (9) Bausch and Lomb (United States); (10) Novartis (United States); (11) Otsuka (United States); (12) Allergan (United States); (13) Allergan (United States); (14) Allergan (United States); (15) Bausch and Lomb (United States); (16) Alcon (United States); (17) Alcon (United States); (18) Advanced Vision Research (United States); (19) Pfizer (United States);

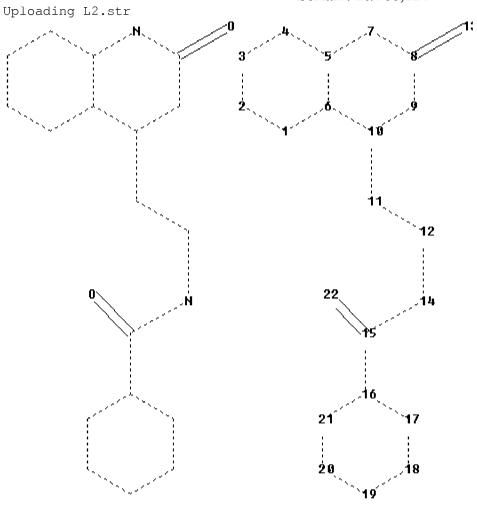
Senju (Japan)

## Serial#: 10/566,214 SEARCH HISTORY

```
FILE 'HCAPLUS' ENTERED AT 17:20:22 ON 20 NOV 2009
              E US2006-566214/APPS
L1
             2 SEA SPE=ON ABB=ON PLU=ON US2006-566214/APPS
    FILE 'REGISTRY' ENTERED AT 17:21:52 ON 20 NOV 2009
               E 2-(4-CHLOROBENZOYLAMINO)-3-(2-OUINOLON-4-YL)PROPIONIC ACID/CN
               E 2-(4-CHLOROBENZOYLAMINO)-3-(2-OUINOLON-4-YL)PROPIONIC ACID/CN
               STRUCTURE UPLOADED
L2.
               D
L3
             8 SEA SSS SAM L2
L4
            99 SEA SSS FUL L2
               STRUCTURE UPLOADED
L5
               D
             4 SEA SUB=L4 SSS SAM L5
1.6
L7
            60 SEA SUB=L4 SSS FUL L5
    FILE 'HCAPLUS' ENTERED AT 17:36:17 ON 20 NOV 2009
L8
           357 SEA SPE=ON ABB=ON PLU=ON L7
    FILE 'REGISTRY' ENTERED AT 17:37:44 ON 20 NOV 2009
               STRUCTURE UPLOADED
L9
               D
            54 SEA SUB=L7 SSS FUL L9
L10
L11
               STRUCTURE UPLOADED
            48 SEA SUB=L7 SSS FUL L11
L12
L13
               STRUCTURE UPLOADED
            35 SEA SUB=L7 SSS FUL L13
L14
    FILE 'HCAPLUS' ENTERED AT 17:45:01 ON 20 NOV 2009
L15
          356 SEA SPE=ON ABB=ON PLU=ON L14
L16
         57021 SEA SPE=ON ABB=ON PLU=ON SALIVA+PFT/CT OR SALIVA?/BI
L17
           11 SEA SPE=ON ABB=ON PLU=ON L16(L)ACCELERATION/OBI
L18
            1 SEA SPE=ON ABB=ON PLU=ON L15 AND L17
           630 SEA SPE=ON ABB=ON PLU=ON XEROSTOMIA+OLD, PFT/CT
L19
             2 SEA SPE=ON ABB=ON PLU=ON L15 AND L19
L20
    FILE 'REGISTRY' ENTERED AT 17:49:15 ON 20 NOV 2009
              E 4-QUINOLINEPROPANOIC ACID, A-"((4-CHLOROBENZOYL)AMINO)"
             1 SEA SPE=ON ABB=ON PLU=ON "4-QUINOLINEPROPANOIC ACID,
L21
               A-((4-CHLOROBENZOYL)AMINO)-1,2-DIHYDRO-2-OXO-"/CN
    FILE 'HCAPLUS' ENTERED AT 17:53:51 ON 20 NOV 2009
L22
          353 SEA SPE=ON ABB=ON PLU=ON L21
           112 SEA SPE=ON ABB=ON PLU=ON NAGAMOTO H?/AU
L23
           78 SEA SPE=ON ABB=ON PLU=ON KOHASHI M?/AU
L24
          1964 SEA SPE=ON ABB=ON PLU=ON OKA H?/AU
L25
L26
            3 SEA SPE=ON ABB=ON PLU=ON L15 AND L16
            3 SEA SPE=ON ABB=ON PLU=ON L18 OR L20 OR L26
L27
            3 SEA SPE=ON ABB=ON PLU=ON L27 AND ((L23 OR L24 OR L25))
L28
            1 SEA SPE=ON ABB=ON PLU=ON L23 AND L24 AND L25
L29
L30
             3 SEA SPE=ON ABB=ON PLU=ON L28 OR L29
    FILE 'REGISTRY' ENTERED AT 17:57:08 ON 20 NOV 2009
               SET SMARTSELECT ON
L31
               SEL PLU=ON L21 1- NAME : 4 TERMS
               SET SMARTSELECT OFF
```

```
FILE 'MEDLINE' ENTERED AT 17:57:08 ON 20 NOV 2009
          228 SEA SPE=ON ABB=ON PLU=ON L31
228 SEA SPE=ON ABB=ON PLU=ON L21 OR L32
26444 SEA SPE=ON ABB=ON PLU=ON SALIVA/CT
L32
L33
L34
L35
          5906 SEA SPE=ON ABB=ON PLU=ON ACCELERATION/CT
L36
             4 SEA SPE=ON ABB=ON PLU=ON L34 AND L35
L37
          3041 SEA SPE=ON ABB=ON PLU=ON XEROSTOMIA/CT
           O SEA SPE=ON ABB=ON PLU=ON L33 AND L36 AND L37
L38
         0 SEA SPE=ON ABB=ON PLU=ON L33 AND L36
1 SEA SPE=ON ABB=ON PLU=ON L33 AND L37
1 SEA SPE=ON ABB=ON PLU=ON L33 AND L34
1 SEA SPE=ON ABB=ON PLU=ON L40 OR L41
8594 SEA SPE=ON ABB=ON PLU=ON "SJOGREN'S SYNDROME"/CT
L39
L40
L41
L42
L43
L44
          3 SEA SPE=ON ABB=ON PLU=ON L33 AND L43
L45
             3 SEA SPE=ON ABB=ON PLU=ON L42 OR L44
           3 SEA SPE=UN ABB=UN PLU=UN L42 UN L47 AND L47 AND
L46
L47
L48
L49
            O SEA SPE=ON ABB=ON PLU=ON L46 AND L47 AND L48
L50
             2 SEA SPE=ON ABB=ON PLU=ON L45 AND ((L46 OR L47 OR L48))
    FILE 'HCAPLUS' ENTERED AT 18:08:01 ON 20 NOV 2009
L51 3703 SEA SPE=ON ABB=ON PLU=ON SJOGREN SYNDROME+OLD, PFT/CT
           4 SEA SPE=ON ABB=ON PLU=ON L15 AND L51
L52
              4 SEA SPE=ON ABB=ON PLU=ON L27 OR L52
L53
    FILE 'REGISTRY' ENTERED AT 18:10:25 ON 20 NOV 2009
                SET SMARTSELECT ON
L54
                SEL PLU=ON L21 1- NAME : 4 TERMS
                SET SMARTSELECT OFF
    FILE 'WPIX, BIOSIS' ENTERED AT 18:10:26 ON 20 NOV 2009
L55 379 SEA SPE=ON ABB=ON PLU=ON L54
           379 SEA SPE=ON ABB=ON PLU=ON L21 OR L55
          3824 SEA SPE=ON ABB=ON PLU=ON SALIVA?(10A)(ACCEL? OR INCREAS?)
           0 SEA SPE=ON ABB=ON PLU=ON L56 AND L57
L58
         0 SEA SPE=ON ABB=ON PLU=ON L56(P) L57
0 SEA SPE=ON ABB=ON PLU=ON L56(30A) L57
4470 SEA SPE=ON ABB=ON PLU=ON XEROSTO? OR DRY(3A) MOUTH?
L59
L60
L61
L62
          O SEA SPE=ON ABB=ON PLU=ON L56 AND L61
L63
             0 SEA SPE=ON ABB=ON PLU=ON L56(30A) L61
            82 SEA SPE=ON ABB=ON PLU=ON NAGAMOTO H?/AU
          73 SEA SPE=ON ABB=ON PLU=ON KOHASHI M?/AU
L65
          3368 SEA SPE=ON ABB=ON PLU=ON OKA H?/AU
1 SEA SPE=ON ABB=ON PLU=ON L64 AND L65 AND L66
L66
L67
    FILE 'REGISTRY' ENTERED AT 18:18:27 ON 20 NOV 2009
                SET SMARTSELECT ON
L68
                 SEL PLU=ON L21 1- NAME : 4 TERMS
                 SET SMARTSELECT OFF
    FILE 'EMBASE' ENTERED AT 18:18:28 ON 20 NOV 2009
L69 381 SEA SPE=ON ABB=ON PLU=ON L68
L70
           381 SEA SPE=ON ABB=ON PLU=ON L21 OR L69
         17167 SEA SPE=ON ABB=ON PLU=ON XEROSTOMIA/CT
          9448 SEA SPE=ON ABB=ON PLU=ON SJOEGREN SYNDROME/CT
          7904 SEA SPE=ON ABB=ON PLU=ON SALIVA/CT
L73
           7 SEA SPE=ON ABB=ON PLU=ON L70 AND L71
12 SEA SPE=ON ABB=ON PLU=ON L70 AND L72
L74
L75 12 SEA SPE=ON ABB=ON FLO-ON L70 AND L73
```

		Serial#: 10/566,214
L77		2 SEA SPE=ON ABB=ON PLU=ON L75 AND MANAGEMENT/TI
L78		5 SEA SPE=ON ABB=ON PLU=ON L75 NOT (EYE OR RHEUMATOLOGIC)/TI
L79		5 SEA SPE=ON ABB=ON PLU=ON L75 AND MANAGEMENT/TI 5 SEA SPE=ON ABB=ON PLU=ON L75 NOT (EYE OR RHEUMATOLOGIC)/TI 7 SEA SPE=ON ABB=ON PLU=ON L77 OR L78 9 SEA SPE=ON ABB=ON PLU=ON L74 OR L79
L80		
L81		35 SEA SPE=ON ABB=ON PLU=ON NAGAMOTO H?/AU
		15 SEA SPE=ON ABB=ON PLU=ON KOHASHI M?/AU
		863 SEA SPE=ON ABB=ON PLU=ON OKA H?/AU
L84 L85		0 SEA SPE=ON ABB=ON PLU=ON L81 AND L82 AND L83 2 SEA SPE=ON ABB=ON PLU=ON L80 AND (L81 OR L82 OR L83)
поэ		2 SEA SPE-ON ADD-ON PRO-ON ROO AND (ROT ON ROZ ON ROS)
	FILE	'REGISTRY' ENTERED AT 18:31:38 ON 20 NOV 2009
	FILE	'HCAPLUS' ENTERED AT 18:31:46 ON 20 NOV 2009 D STAT QUE L30
	FILE	'MEDLINE' ENTERED AT 18:32:00 ON 20 NOV 2009 D STAT QUE L50
	FILE	'WPIX, BIOSIS' ENTERED AT 18:32:54 ON 20 NOV 2009 D STAT QUE L67
	FILE	'EMBASE' ENTERED AT 18:33:02 ON 20 NOV 2009 D STAT QUE L85
L86	FILE	'HCAPLUS, MEDLINE, WPIX, EMBASE' ENTERED AT 18:33:28 ON 20 NOV 2009 3 DUP REMOVE L30 L50 L67 L85 (5 DUPLICATES REMOVED) ANSWERS '1-3' FROM FILE HCAPLUS D L86 IBIB ABS HITSTR 1-3
	FILE	'HCAPLUS' ENTERED AT 18:33:53 ON 20 NOV 2009 D STAT QUE L53
	FILE	'MEDLINE' ENTERED AT 18:34:11 ON 20 NOV 2009
	FILE	'HCAPLUS' ENTERED AT 18:34:19 ON 20 NOV 2009
L87		1 SEA SPE=ON ABB=ON PLU=ON L53 NOT L30
	FILE	'MEDLINE' ENTERED AT 18:34:42 ON 20 NOV 2009
L88		D STAT QUE L45 1 SEA SPE=ON ABB=ON PLU=ON L45 NOT L50
гоо		I SEA SPE-ON ABB-ON PEO-ON E45 NOT E50
	FILE	'WPIX, BIOSIS' ENTERED AT 18:35:23 ON 20 NOV 2009  D STAT QUE L60 D STAT QUE L62
	FILE	'EMBASE' ENTERED AT 18:35:44 ON 20 NOV 2009
L89		D STAT QUE L80 7 SEA SPE=ON ABB=ON PLU=ON L80 NOT L85
L90	FILE	'HCAPLUS, MEDLINE, EMBASE' ENTERED AT 18:36:18 ON 20 NOV 2009  8 DUP REMOVE L87 L88 L89 (1 DUPLICATE REMOVED)  ANSWER '1' FROM FILE HCAPLUS  ANSWER '2' FROM FILE MEDLINE  ANSWERS '3-8' FROM FILE EMBASE



```
chain nodes :
11  12  13  14  15  22
ring nodes :
1  2  3  4  5  6  7  8  9  10  16  17  18  19  20  21
chain bonds :
8-13  10-11  11-12  12-14  14-15  15-16  15-22
ring bonds :
1-2  1-6  2-3  3-4  4-5  5-6  5-7  6-10  7-8  8-9  9-10  16-17  16-21  17-18  18-19
19-20  20-21
exact/norm bonds :
1-2  1-6  2-3  3-4  4-5  5-6  5-7  6-10  7-8  8-9  8-13  9-10  10-11  11-12  12-14
14-15  15-16  15-22  16-17  16-21  17-18  18-19  19-20  20-21
```

# Match level:

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:CLASS

Uploading L5.str

```
chain nodes :
11 12 13 14 15 22 23 24 25 26
ring nodes :
1 2 3 4 5 6 7 8 9 10 16 17 18 19 20 21
chain bonds :
8-13 \quad 10-11 \quad 11-12 \quad 12-14 \quad 12-23 \quad 14-15 \quad 15-16 \quad 15-22 \quad 19-26 \quad 23-24 \quad 23-25
ring bonds :
1-2 \quad 1-6 \quad 2-3 \quad 3-4 \quad 4-5 \quad 5-6 \quad 5-7 \quad 6-10 \quad 7-8 \quad 8-9 \quad 9-10 \quad 16-17 \quad 16-21 \quad 17-18 \quad 18-19
19-20 20-21
exact/norm bonds :
5-7 6-10 7-8 8-9 8-13 9-10 12-14 14-15 15-22 23-24 23-25
exact bonds :
10-11 11-12 12-23 15-16 19-26
normalized bonds :
1-2 \quad 1-6 \quad 2-3 \quad 3-4 \quad 4-5 \quad 5-6 \quad 16-17 \quad 16-21 \quad 17-18 \quad 18-19 \quad 19-20 \quad 20-21
Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:Atom 17:Atom 18:Atom 19:Atom
20:Atom 21:Atom
22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS
```

```
chain nodes :
11  12  13  14  15  22  23  24  25  26
ring nodes :
1  2  3  4  5  6  7  8  9  10  16  17  18  19  20  21
chain bonds :
8-13  10-11  11-12  12-14  12-23  14-15  15-16  15-22  19-26  23-24  23-25
ring bonds :
1-2  1-6  2-3  3-4  4-5  5-6  5-7  6-10  7-8  8-9  9-10  16-17  16-21  17-18  18-19
19-20  20-21
exact/norm bonds :
5-7  6-10  7-8  8-9  8-13  9-10  12-14  14-15  15-22
exact bonds :
10-11  11-12  12-23  15-16  19-26
normalized bonds :
1-2  1-6  2-3  3-4  4-5  5-6  16-17  16-21  17-18  18-19  19-20  20-21  23-24  23-25
```

12:3 E exact RC ring/chain

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:Atom 17:Atom 18:Atom 19:Atom

20:Atom 21:Atom

22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS